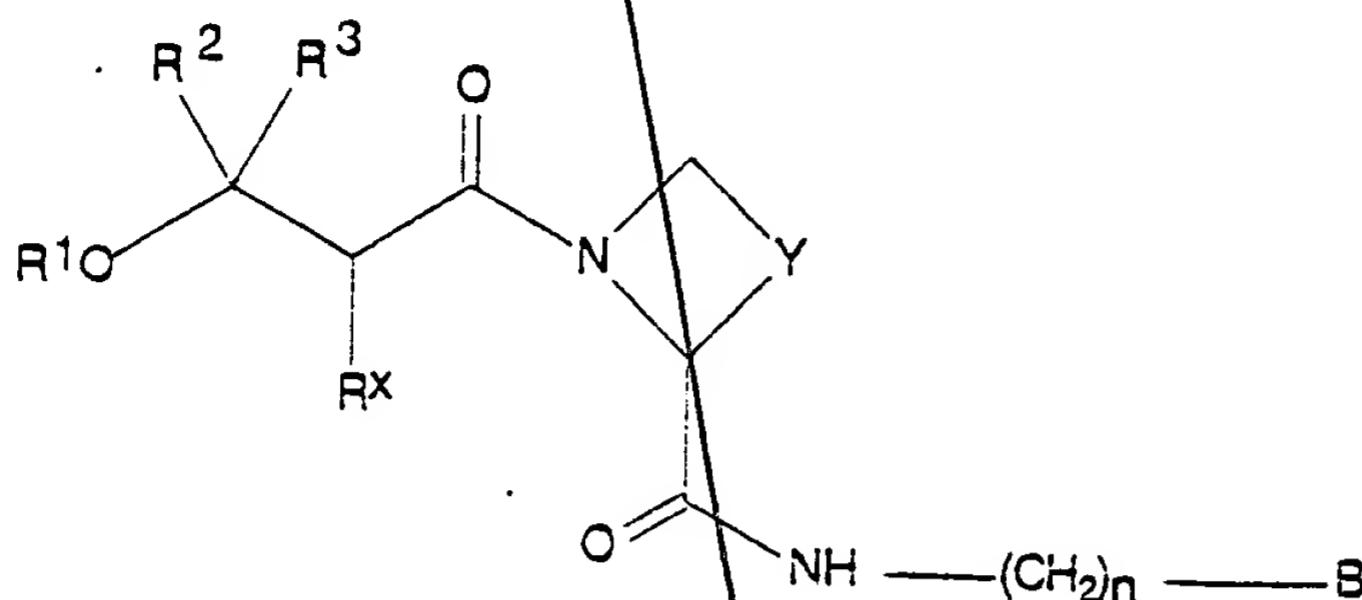


1. (Amended) A compound of formula I,



wherein

R¹ represents H, C(O)R¹¹, SiR¹²R¹³R¹⁴ or C₁₋₆ alkyl which latter group is optionally substituted or terminated by one or more substituent selected from OR¹⁵ or (CH₂)_qR¹⁶;

R¹², R¹³ and R¹⁴ independently represent H, phenyl or C₁₋₆ alkyl;

R¹⁶ represents C₁₋₄ alkyl, phenyl, OH, C(O)OR¹⁷ or C(O)N(H)R¹⁸;

R¹⁸ represents H, C₁₋₄ alkyl or CH₂C(O)OR¹⁹;

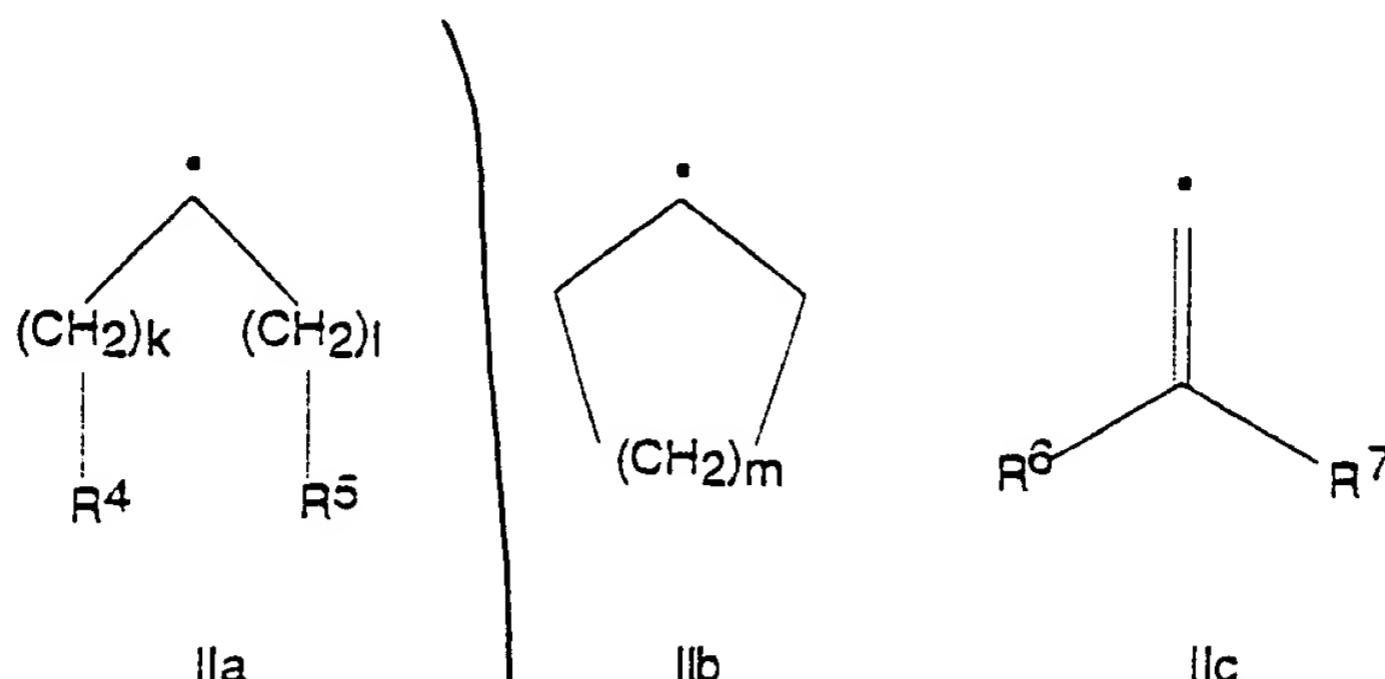
R¹⁵ and R¹⁷ independently represent H, C₁₋₆ alkyl or C₇₋₉ alkylphenyl;

R¹¹ and R¹⁹ independently represent H or C₁₋₄ alkyl; and

q represents 0, 1 or 2;

R² and R³ independently represent H, C₁₋₄ alkyl, cyclohexyl or phenyl;

R^x represents a structural fragment of formula IIa, IIb or IIc,



wherein

k, l and m independently represent 0, 1, 2, 3 or 4;

R^4 and R^5 independently represent H, $Si(Me)_3$, 1- or 2-naphthyl, a polycyclic hydrocarbyl group, $CHR^{41}R^{42}$ or $C_{1.4}$ alkyl (which latter group is optionally substituted by one or more fluorine atoms), or $C_{3.8}$ cycloalkyl phenyl, methylenedioxyphenyl, benzodioxanyl, benzofuranyl, dihydrobenzofuranyl, benzothiazolyl, benzoxazolyl, benzimidazolyl, coumaranonyl, coumarinyl or dihydrocoumarinyl (which latter twelve groups are optionally substituted by one or more of $C_{1.4}$ alkyl (which latter group is optionally substituted by one or more halo substituent), $C_{1.4}$ alkoxy, halo, hydroxy, cyano, nitro, SO_2NH_2 , $C(O)OH$ or $N(H)R^{43}$);

R⁴¹ and R⁴² independently represent cyclohexyl or phenyl;

R^6 and R^7 independently represent H, C_{1-4} alkyl, C_{3-8} cycloalkyl, phenyl (which latter group is are optionally substituted by one or more of C_{1-4} alkyl (which latter group is optionally substituted by one or more halo substituent).

$C_{1.4}$ alkoxy, halo, hydroxy, cyano, nitro, SO_2NH_2 , $C(O)OH$ or $N(H)R^{44}$) or together with the carbon atom to which they are attached form a $C_{3.8}$ cycloalkyl ring;

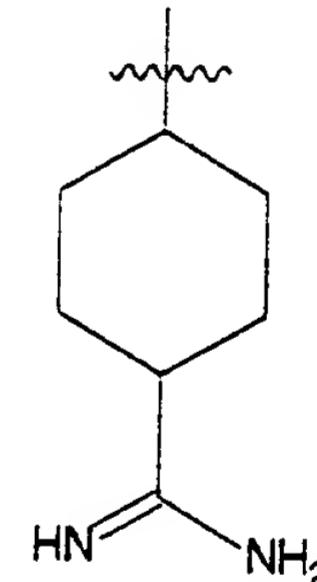
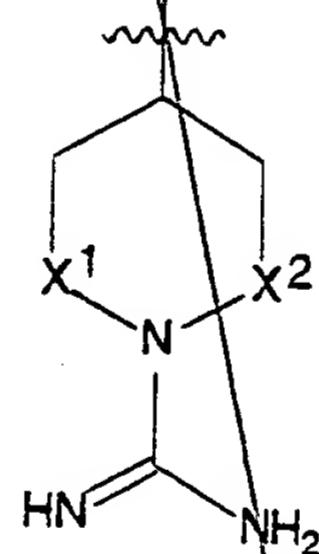
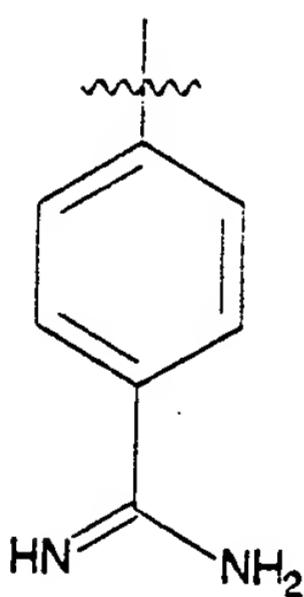
R^{43} and R^{44} independently represent H or $C(O)R^{45}$; and

R^{45} represents H, $C_{1.4}$ alkyl or $C_{1.4}$ alkoxy;

Y represents $(CH_2)_2$, $CH=CH$, $(CH_2)_3$, $CH_2CH=CH$ or $CH=CHCH_2$, which latter three groups are optionally substituted by $C_{1.4}$ alkyl, methylene, oxo or hydroxy;

n represents 0, 1, 2, 3 or 4; and

B represents a structural fragment of formula IVa, IVb or IVc



wherein

X^1 and X^2 independently represents a single bond or CH_2 ;

or a pharmaceutically acceptable salt thereof.

A²
3. (Amended) A compound of formula I, as defined in Claim 1, wherein
 R^1 represents optionally substituted C_{1-6} alkyl or H.

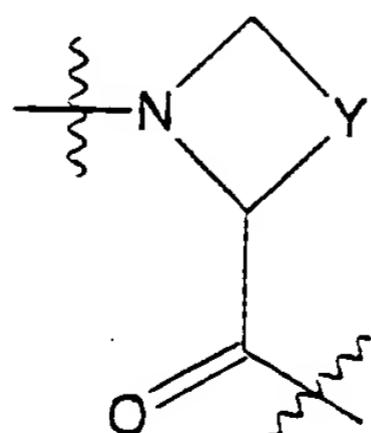
5. (Amended) A compound of formula I, as defined in claim 1, wherein
 R^x represents a structural fragment of formula IIa.

6. (Amended) A compound of formula I, as defined in claim 1, wherein
Y represents $(CH_2)_2$.

7. (Amended) A compound of formula I, as defined in Claim 1, wherein
n represents 1.

8. (Amended) A compound of formula I, as defined in Claim 1, wherein
B represents a structural fragment of formula IVa..

9. (Amended) A compound of formula I, as defined in claim 1, wherein
the fragment



is in the S-configuration.

10. (Amended) A compound as claimed in Claim 1 which is

(*R,S*)-PhCH(CH₂OH)-C(O)-Pro-(*R,S*)-Hig;
(*S*)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R,S*)-3-aminophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-3-(methylamino)phenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-3-(methylamino)phenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-PhCH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-3-(trifluoromethyl)phenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-3-(trifluoromethyl)phenyl-CH(CH₂OH)-C(O)-Pro-Pab;
3 (*R,S*)-3-hydroxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-((3-chloro-5-methylphenyl)-CH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-((3-chloro-5-methylphenyl)-CH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-3-fluorophenyl-CH(CH₂OH)CO-Pro-Pab;
(*R*)-3-fluorophenyl-CH(CH₂OH)CO-Pro-Pab;
(*S*)-3-chlorophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-3-chlorophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R,S*)-3,5-dimethylphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-3,5-bis(trifluoromethyl)phenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-3,5-bis(trifluoromethyl)phenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R,S*)-3-methoxy-5-methylphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(*R,S*)-(2,5-dimethoxyphenyl)-CH(CH₂OH)-C(O)-Pro-Pab;
(*R,S*)-(3,5-dimethoxyphenyl)-CH(CH₂OH)-C(O)-Pro-Pab;
(*R,S*)-3,4-(methylenedioxyphenyl)-CH(CH₂OH)-C(O)-Pro-Pab;
(*S*)-3-(2-naphthyl)-CH(CH₂OH)-C(O)-Pro-Pab;
(*R*)-3-(2-naphthyl)-CH(CH₂OH)-C(O)-Pro-Pab;

(R)-2,5-dimethylphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(S)-2,5-dimethylphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(R)-3-methoxy-4-hydroxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(S)-3-methoxy-4-hydroxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(R)-3,5-dichlorophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(S)-3,5-dichlorophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(R)-2,3-dimethoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(S)-2,3-dimethoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(R)-3-methoxy-5-chlorophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(S)-3-methoxy-5-chlorophenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(R)-2-methyl-5-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(S)-2-methyl-5-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab;
(R,S)-Ph-C(Me)(CH₂OMe)-C(O)-Pro-Pab;
(R)-2,3-(methylenedioxyphenyl)-CH(CH₂OH)-C(O)-Pro-Pab; or
(S)-2,3-(methylenedioxyphenyl)-CH(CH₂OH)-C(O)-Pro-Pab
or a pharmaceutically acceptable salt thereof.

19. (Amended) A compound as claimed in Claim 17 which is

(R,S)-Ph-CH(CH₂OH)-C(O)-Pro-Pab-OH;
(S)-3-methoxyphenyl-CH(CH₂OH)CO-Pro-Pab(Z);
(R)-3-methoxyphenyl-CH(CH₂OH)CO-Pro-Pab(Z);
(S)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab-OH;
(R)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab-OH;
(S)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab-OC(O)Et;
(R)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab-OC(O)Et;
(S)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab-OC(O)CH₃;

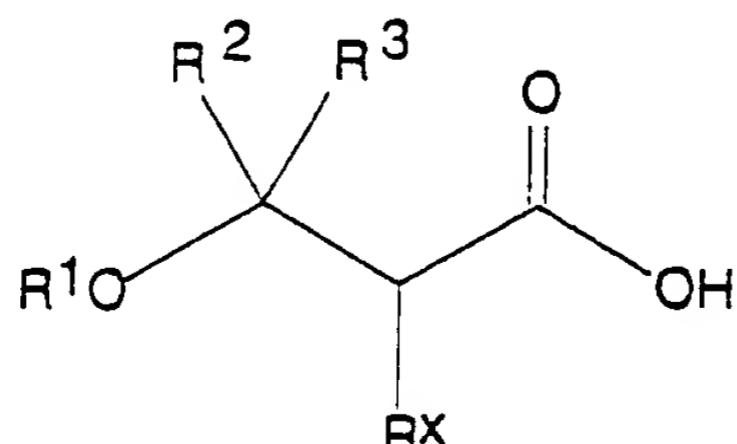
(R)-3-methoxyphenyl-CH(CH₂OH)-C(O)-Pro-Pab-OC(O)CH₃;
(R,S)-3-Ph-C(Me)(CH₂OMe)-C(O)-Pro-Pab(Z); or
(R,S)-3-methylphenyl-CH(CH₂OAc)-C(O)-Pro-Pab-OMe;
or a pharmaceutically acceptable salt thereof.

11/10/01
20. (Amended) A pharmaceutical formulation including a compound as defined in claim 1, or a pharmaceutically acceptable salt thereof, in admixture with a pharmaceutically acceptable adjuvant, diluent or carrier..

11/10/01
28. (Amended) A method of treatment of a condition where inhibition of thrombin is required which method comprises administration of a therapeutically effective amount of a compound as defined in claim 1, or a pharmaceutically acceptable salt thereof, to a person suffering from, or susceptible to, such a condition.

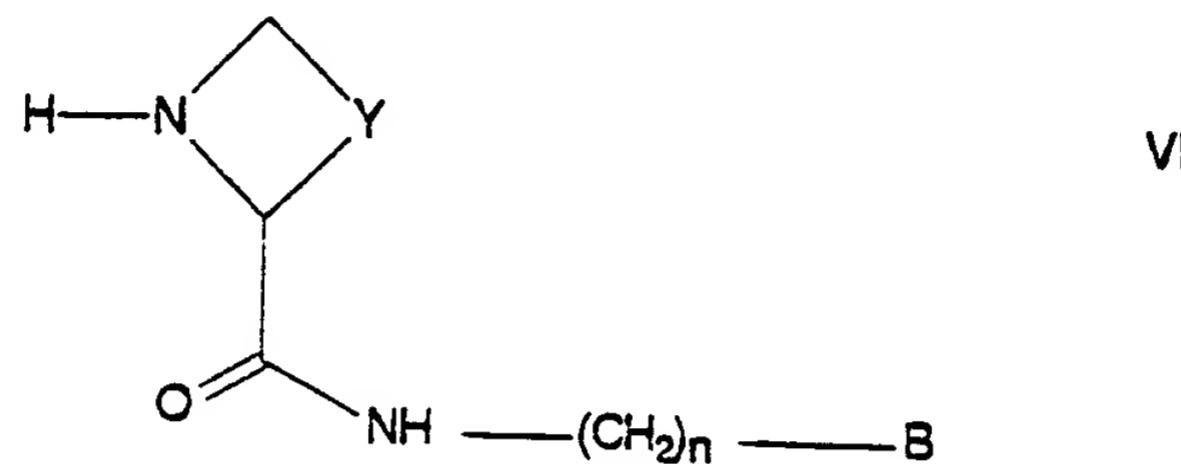
11/10/01
32. (Amended) A process for the preparation of compounds of formula I as defined in claim 1, which comprises:

(a) the coupling of a compound of formula V,



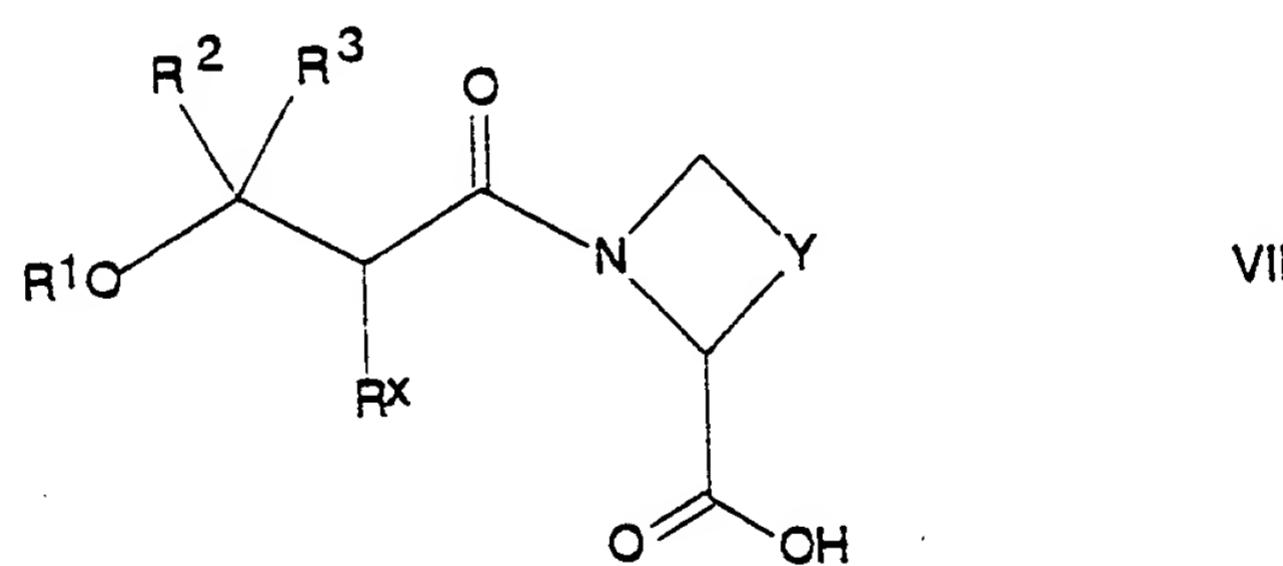
V

wherein R^1 , R^2 R^3 and R^x are as defined in Claim 1, with a compound of formula VI,



wherein Y, n and B are as defined in Claim 1; or

(b) the coupling of a compound of formula VII,



wherein R^1 , R^2 , R^3 , R^x and Y are as defined in Claim 1 with a compound of formula VIII,

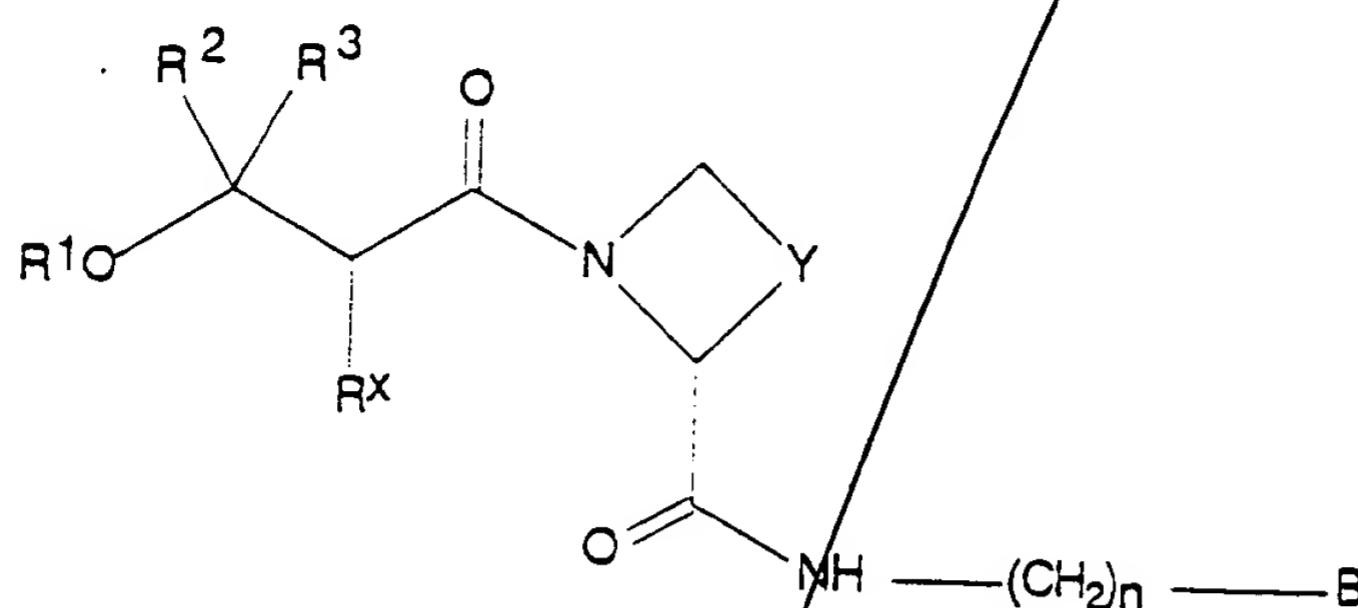


wherein n and B are as defined in Claim 1.

Please add the following new claim:

33. (New) A compound of formula I, as defined in Claim 1, wherein R² and R³ are both H.

1. (Twice Amended) A compound of formula I,



wherein

R^1 represents H , $\text{C}(\text{O})\text{R}^{11}$, $\text{SiR}^{12}\text{R}^{13}\text{R}^{14}$ or C_{1-6} alkyl which latter group is optionally substituted or terminated by one or more substituent selected from the group consisting of OR^{15} and $(\text{CH}_2)_q\text{R}^{16}$;

R^{12} , R^{13} and R^{14} independently represent H , phenyl or C_{1-6} alkyl;

R^{16} represents C_{1-4} alkyl, phenyl, OH , $\text{C}(\text{O})\text{OR}^{17}$ or $\text{C}(\text{O})\text{N}(\text{H})\text{R}^{18}$;

R^{18} represents H , C_{1-4} alkyl or $\text{CH}_2\text{C}(\text{O})\text{OR}^{19}$;

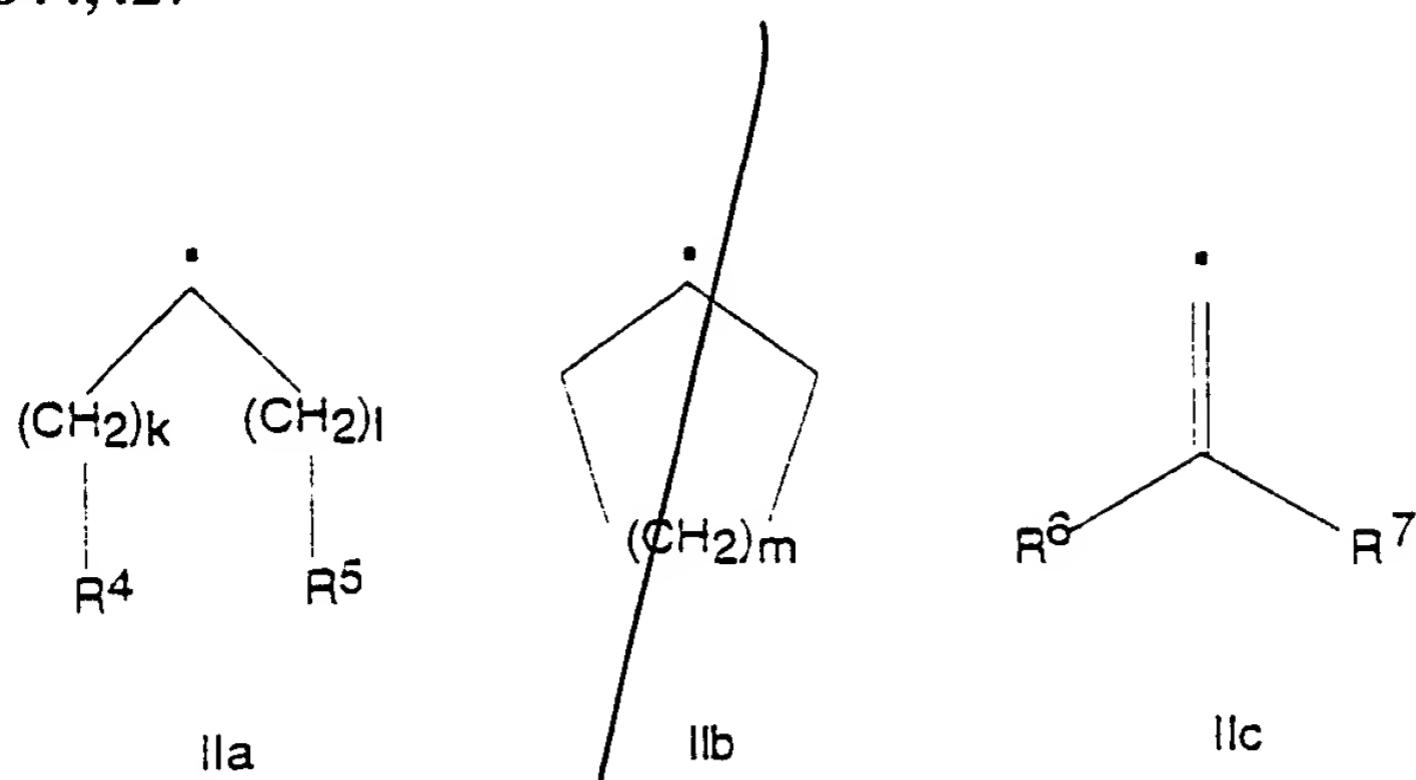
R^{15} and R^{17} independently represent H , C_{1-6} alkyl or C_{7-9} alkylphenyl;

R^{11} and R^{19} independently represent H or C_{1-4} alkyl; and

q represents 0, 1 or 2;

R^2 and R^3 are both hydrogen;

R^x represents a structural fragment of formula IIa, IIb or IIc,



wherein

k, l and m independently represent 0, 1, 2, 3 or 4;

R^4 and R^5 independently represent H, $Si(Me)_3$, 1- or 2-naphthyl, a polycyclic hydrocarbyl group, $CHR^{41}R^{42}$ or C_{1-4} alkyl (which latter group is optionally substituted by one or more fluorine atoms), or C_{3-8} cycloalkyl, phenyl, methylenedioxophenyl, benzodioxanyl, benzofuranyl, dihydrobenzofuranyl, benzothiazolyl, benzoxazolyl, benzimidazolyl, coumaranonyl, coumarinyl or dihydrocoumarinyl (which latter twelve groups are optionally substituted by one or more of C_{1-4} alkyl (which latter group is optionally substituted by one or more halo substituent), C_{1-4} alkoxy, halo, hydroxy, cyano, nitro, SO_2NH_2 , $C(O)OH$ or $N(H)R^{43}$).

R^{41} and R^{42} independently represent cyclohexyl or phenyl;

R^6 and R^7 independently represent H, C₁₋₄ alkyl, C₃₋₈ cycloalkyl, phenyl (which latter group is are optionally substituted by one or more of C₁₋₄ alkyl (which latter group is optionally substituted by one or more halo substituent), C₁₋₄ alkoxy, halo, hydroxy, cyano, nitro, SO₂NH₂, C(O)OH or N(H)R⁴⁴) or together

with the carbon atom to which they are attached form a C₃₋₈ cycloalkyl ring;

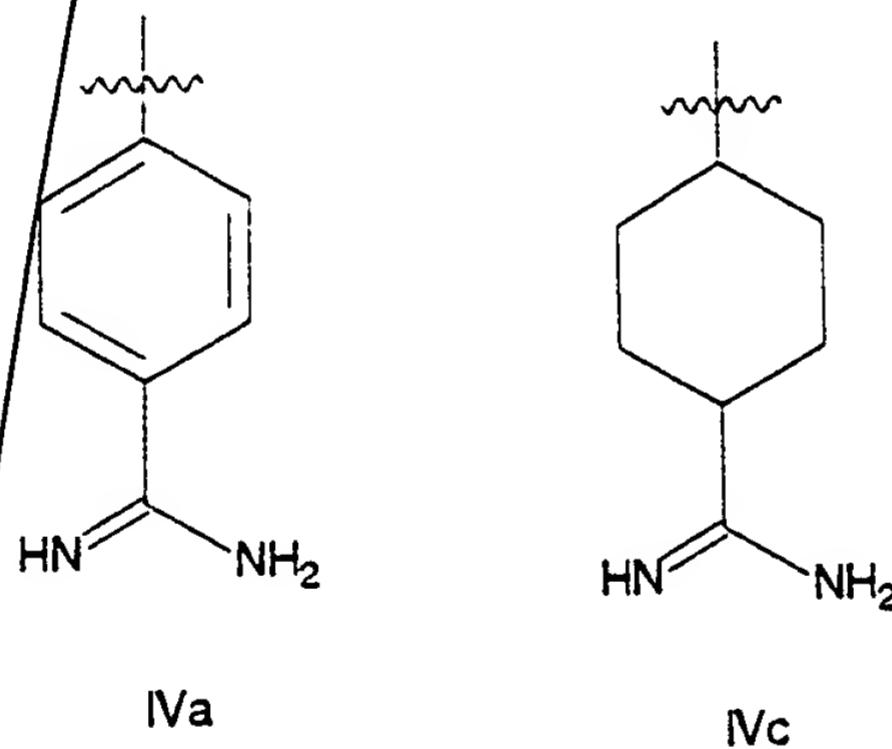
R⁴³ and R⁴⁴ independently represent H or C(O)R⁴⁵; and

R⁴⁵ represents H, C₁₋₄ alkyl or C₁₋₄ alkoxy;

Y represents (CH₂)₂, CH=CH, (CH₂)₃, CH₂CH=CH or CH=CHCH₂, which latter three groups are optionally substituted by C₁₋₄ alkyl, methylene, oxo or hydroxy;

n represents 0, 1, 2, 3 or 4; and

B represents a structural fragment of formula IVa or IVc



or a pharmaceutically acceptable salt thereof.

C2
8. (Twice Amended) A compound of formula I, as defined in Claim 1, wherein B represents a structural fragment of formula IVa.

11. (Amended) A compound of formula I, as defined in Claim 1, provided that when R^x represents a structural fragment of formula IIa, then R⁴ and/or R⁵ do/does not represent phenyl substituted by halo-substituted C₁₋₆ alkyl.

12. (Amended) A compound of formula I, as defined in Claim 1, provided that when R^x represents a structural fragment of formula IIa, then R⁴ and/or R⁵ do/does not represent methylenedioxophenyl, benzodioxanyl, benzofuranyl, dihydrobenzofuranyl, benzothiazolyl, benzoxazolyl, benzimidazolyl, coumaranonyl, coumarinyl or dihydrocoumarinyl.

13. (Amended) A compound of formula I, as defined in Claim 1, provided that when R^x represents a structural fragment of formula IIc, then R⁶ and/or R⁷ represent(s) unsubstituted phenyl.

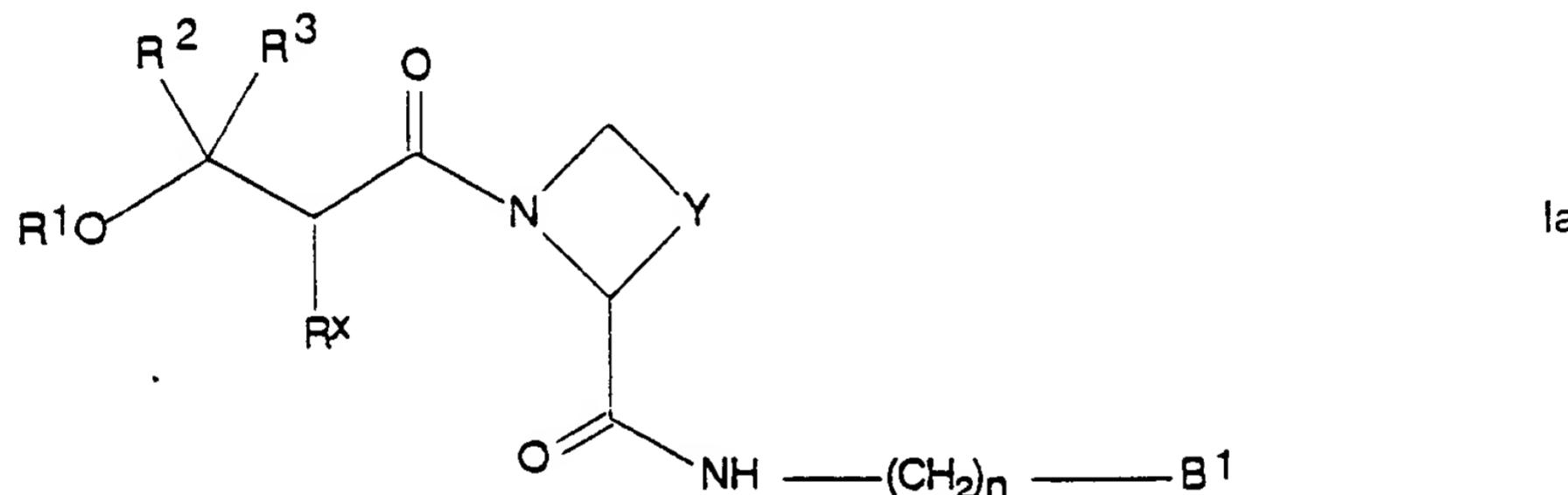
14. (Amended) A compound of formula I, as defined in Claim 1, wherein, when R^x represents a structural fragment of formula IIa, then R⁴ and/or R⁵ represent(s) phenyl substituted by halo-substituted C₁₋₆ alkyl.

15. (Amended) A compound of formula I, as defined in Claim 1, wherein, when R^x represents a structural fragment of formula IIa, then R⁴ and/or R⁵ represent(s) methylenedioxophenyl, benzodioxanyl, benzofuranyl,

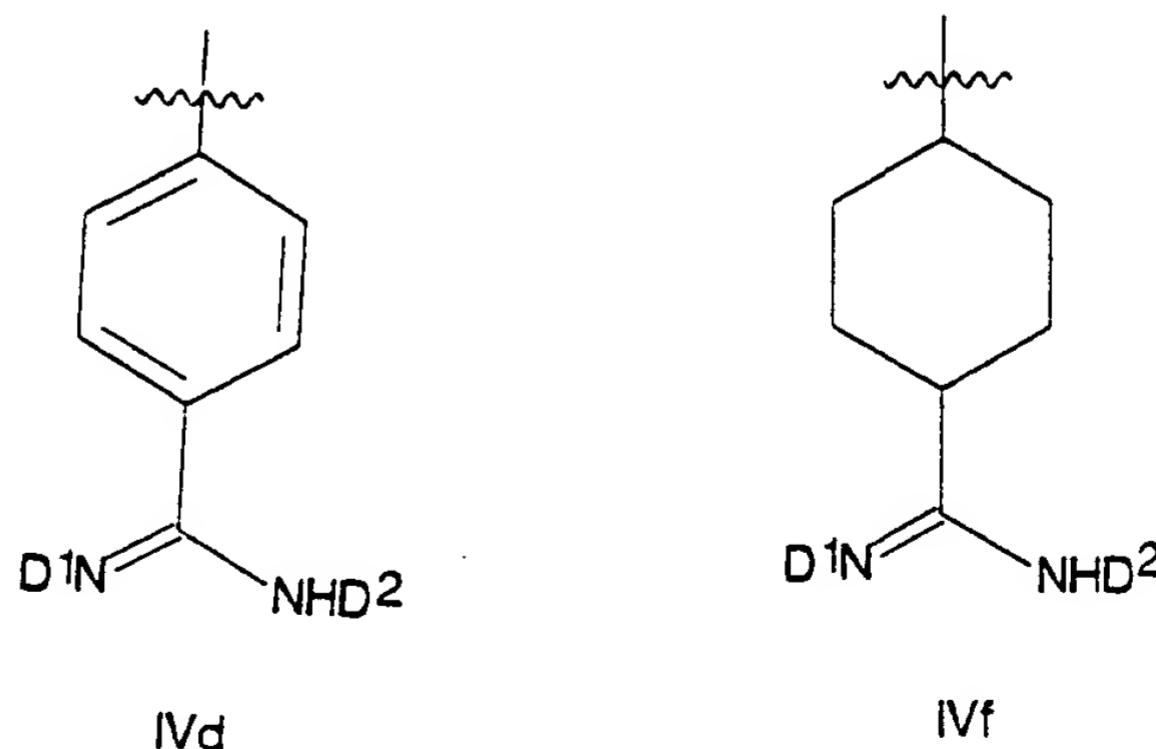
dihydrobenzofuranyl, benzothiazolyl, benzoxazolyl, benzimidazolyl, coumaranonyl, coumarinyl or dihydrocoumarinyl.

16. (Amended) A compound of formula I, as defined in Claim 1, wherein, when R^x represents a structural fragment of formula IIc, then R^6 and/or R^7 represent(s) substituted phenyl.

17. (Amended) A compound of formula Ia,



wherein B^1 represents a structural fragment of formula IVd or IVf



wherein D^1 and D^2 independently represent H, OH, OR^a , OC(O)R^b ,

OC(O)OR^c, C(O)OR^d, or C(O)R^e and R^a, R^b, R^c, R^d and R^e independently represent phenyl, benzyl, (CH₂)₂OC(O)CH₃ or C₁₋₆ alkyl which latter group is optionally interrupted by oxygen; and R¹, R², R³, R^x, Y and n are as defined in

Claim 1, or a pharmaceutically acceptable salt thereof, provided that D¹ and D² do not both represent H.

C4
20. (Twice Amended) A pharmaceutical formulation including a compound as defined in claim 1, or a pharmaceutically acceptable salt thereof, in admixture with a pharmaceutically acceptable adjuvant, diluent or carrier.